U.S. DEPARTMENT OF COMMERCE
Patent and Trademark Office

SEARCH REQUEST FORM

Requestor's Name: B. Dent Serial Number: Dent

Date: 10/26/05 Phone: 308 4544 Art Unit: 1603

Search T
Please wri
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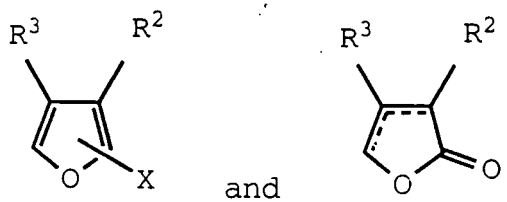
a copy of

-- 35. A compound of a formula selected from

any terms
use attach



and

wherein X is H or hydroxyl;

wherein R^2 is a substituent selected from pyridyl, and aryl optionally substituted at a substitutable position with a radical selected from halo, lower alkyl, lower alkoxy, lower alkylthio, lower alkylsulfinyl, lower alkylsulfonyl, nitro, amino, lower alkylamino, sulfamyl and lower alkylsulfonylamino; and

wherein R^3 is a substituent selected from pyridyl, and aryl optionally substituted at a substitutable position with a radical selected from halo, lower alkyl, lower alkoxy, lower alkylthio, lower alkylsulfinyl, lower alkylsulfonyl, nitro, amino, lower alkylamino, amide, lower alkylsulfonylamino and sulfamyl;

provided that at least one of said R^2 and R^3 substituents is substituted with lower alkylsulfonyl or sulfamyl;

Structure search attached

Ser. No. 08/425022
Applicant Berkstresser

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=> FIL REG
FILE 'REGISTRY' ENTERED AT 11:08:24 ON 30 OCT 95
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
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STRUCTURE FILE UPDATES:  27 OCT 95   HIGHEST RN 169435-71-6
DICTIONARY FILE UPDATES: 29 OCT 95   HIGHEST RN 169435-71-6

TSCA INFORMATION NOW CURRENT THROUGH JUNE 1995

  Please note that search-term pricing does apply when
  conducting SmartSELECT searches.
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this is the search for cpds. wherein X is OH

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=> D QUE L15
L9                  STR
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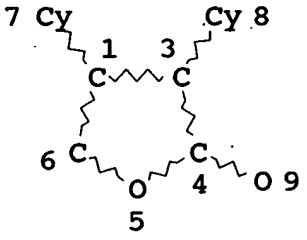


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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS   8

STEREO ATTRIBUTES: NONE
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L13                 STR
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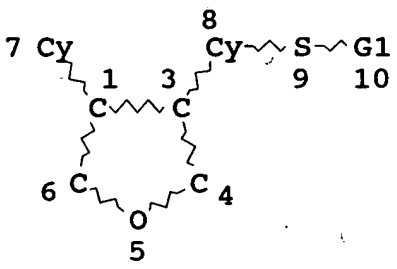


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STEREO ATTRIBUTES: NONE
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=> FIL CAPLUS
FILE 'CAPLUS' ENTERED AT 11:08:31 ON 30 OCT 95
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COPYRIGHT (C) 1995 AMERICAN CHEMICAL SOCIETY (ACS)

FILE COVERS 1967 - 30 Oct 1995  VOL 123 ISS 18
FILE LAST UPDATED: 28 Oct 1995  (951028/ED)

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 HCAplus file when using the FSEARCH command or when conducting
 SmartSELECT searches with large numbers of terms.


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=> S L15
L17             5 L15
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=> D BIB ABS HITSTR 1

L17 ANSWER 1 OF 5 CAPLUS COPYRIGHT 1995 ACS
AN 1995:863354 CAPLUS
TI Substituted spiro compounds for the treatment of inflammation
IN Reitz, David B.; Manning, Robert E.; Huang, Horng Chi; Li, Jinglin
PA G.D. Searle and Co., USA
SO U.S., 40 pp.
CODEN: USXXAM
PI US 5393790 A 950228
AI US 94-194762 940210
DT Patent
LA English
AB A class of substituted spiro compds. is described for use in treating inflammation and inflammation-related disorders. Compds. of particular interest are defined by formula I wherein A is selected from II-V and wherein each of R1 through R10 is independently selected from hydrido, halo, alkyl, alkoxy, alkylthio, cyano, haloalkyl, haloalkoxy, hydroxyalkyl, alkoxyalkyl, hydroxyl, mercapto, alkylsulfonyl, haloalkylsulfonyl and sulfamyl; and wherein n is a no. selected from 0, 1, 2 and 3; or a pharmaceutically-acceptable salt thereof. Thus, e.g., 5-(4-fluorophenyl)-6-[(4-methylsulfonyl)phenyl]spiro[2.4]hept-5-ene (VI), prepd. in 86% yield by cyclization of 1-[2-(4-fluorophenyl)-4,4-di(tosyloxymethyl)cyclopenten-1-yl]-4-(methylsulfonyl)benzene (prepn. given) in presence of NaI/Zn, demonstrated 32% inhibition of rat paw edema and 15% inhibition of the hyperalgesic foot withdrawal at 10 mg/kg body wt., and ID50 of <0.1 .mu.M for inhibition of cyclooxygenase II (ID50 = 14 for COX I).
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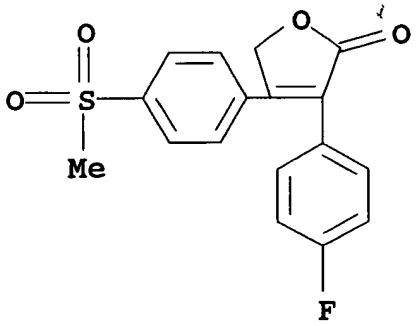
RN 157672-00-9 CAPLUS

CN 2(5H)-Furanone, 3-(4-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



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Ducharme

L17 ANSWER 2 OF 5 CAPLUS COPYRIGHT 1995 ACS
AN 1995:468615 CAPLUS 08/179467 US 5474995 12/12/95
TI (Phenyl)heterocyclic compounds as cyclooxygenase inhibitors
IN Ducharme, Yves; Gauthier, Jacques Yves; Prasit, Petpiboon; Leblanc, Yves; Wang, Zhaoyin; Leger, Serge; Therien, Michel
PA Merck Frosst Canada Inc., Can.
SO PCT Int. Appl., 167 pp.
CODEN: PIXXD2
PI WO 9500501 A2 950105
DS W: AU, BB, BG, BR, BY, CA, CN, CZ, FI, GE, HU, JP, KE, KG, KR, KZ, LK, LV, MD, MG, MN, MW, NO, NZ, PL, RO, RU, SD, SI, SK, TJ, TT, UA, US, UZ
RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG
AI WO 94-CA318 940609
PRAI US 93-82196 930624
US 94-179467 940110
DT Patent
LA English
AB The title compds. I (XYZ form heterocyclic ring; R1 = alkylsulfonyl, aminosulfonyl, etc.; R2 = alkyl, halo, etc.) were disclosed as cyclooxygenase inhibitors (inflammation inhibitors). Example compds. are 2,3-diphenylthiophenes and 3,4-diphenylfuran derivs. Prepd. example compds. are 2-[4-(4-fluorophenyl)-3-thienyl]benzenesulfonamide (II) and 3-(3,4-dichlorophenyl)-4-[4-(methylsulfonyl)phenyl]dihydro-2(3H)-furanone (III).
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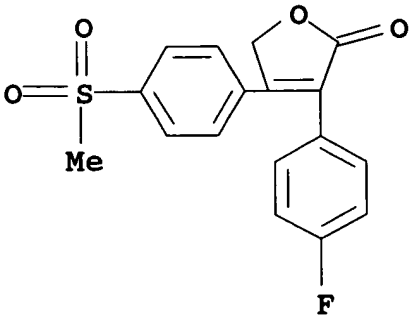

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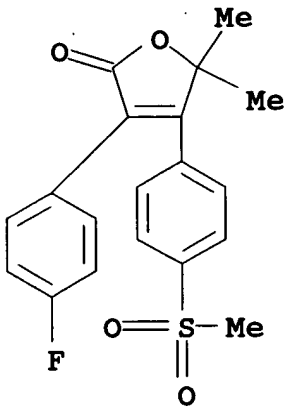

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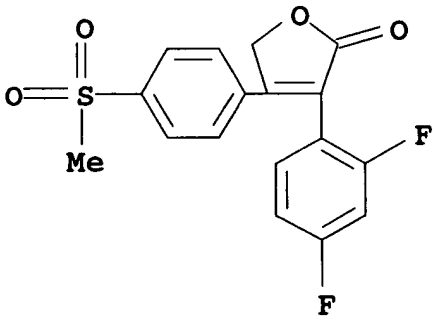

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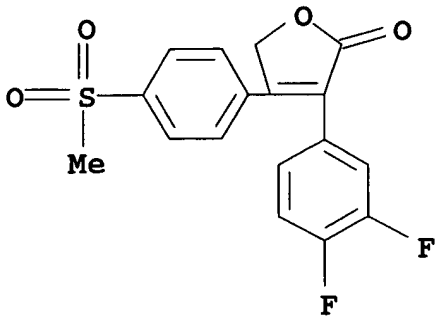

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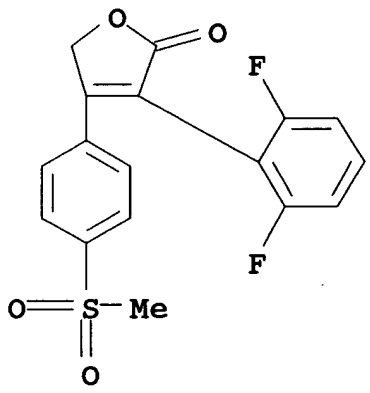

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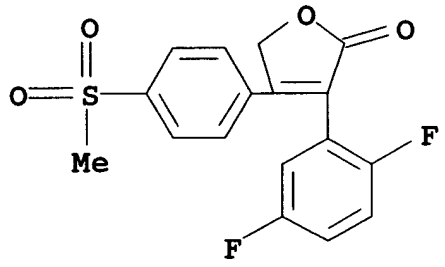

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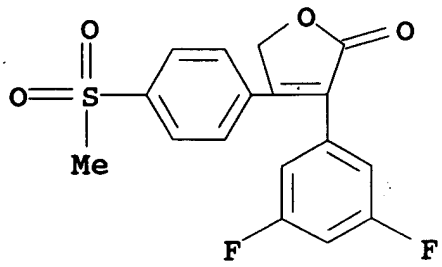

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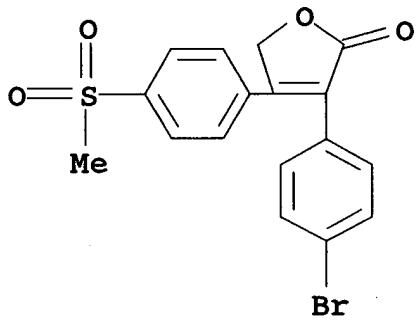

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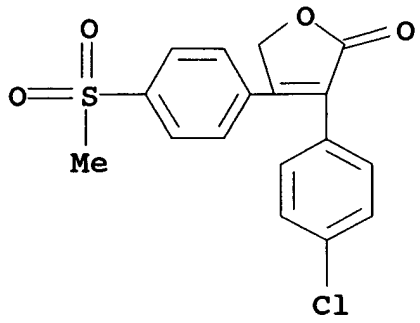

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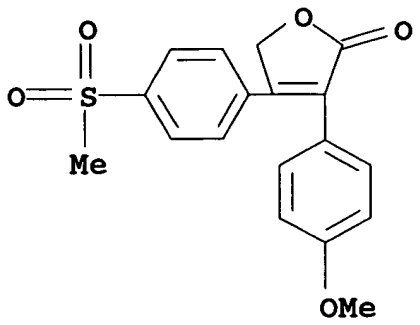

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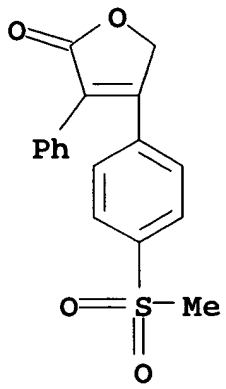

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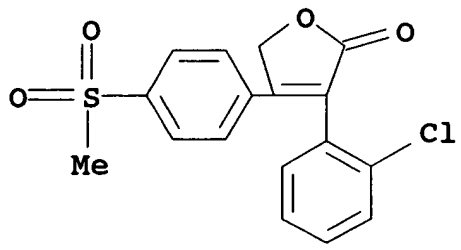

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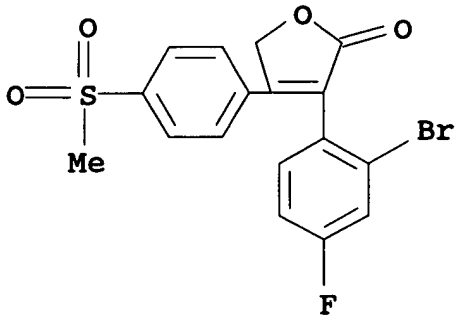

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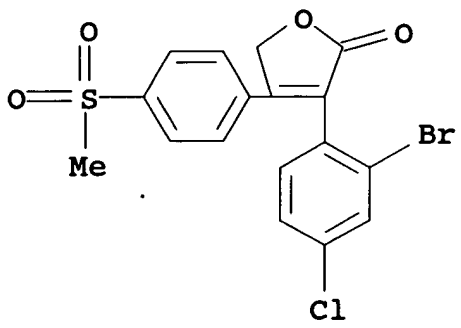

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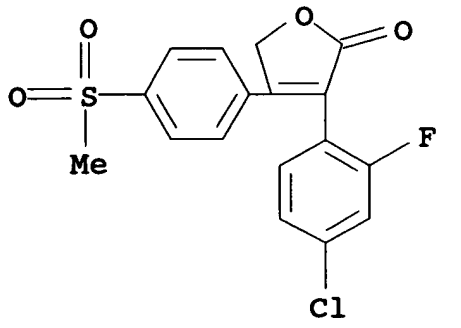

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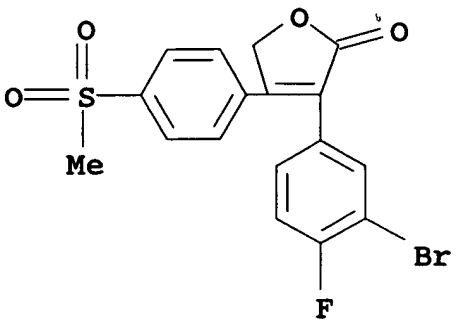

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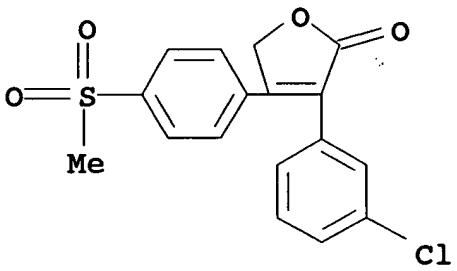

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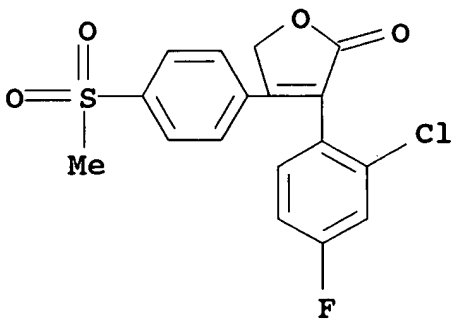

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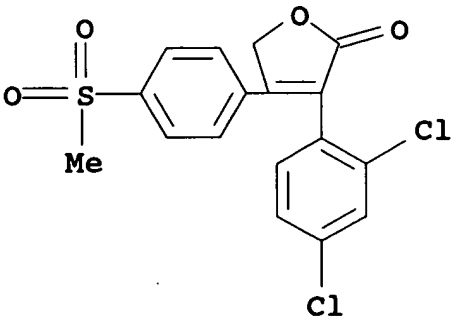
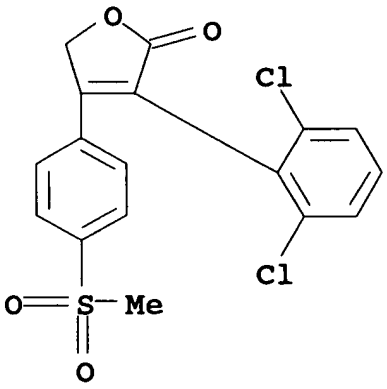
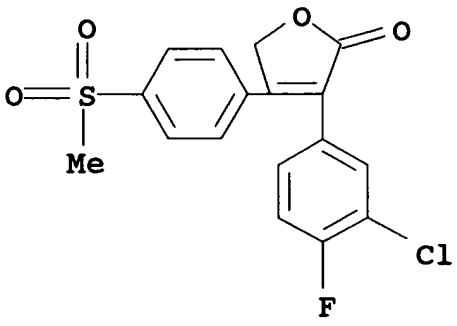


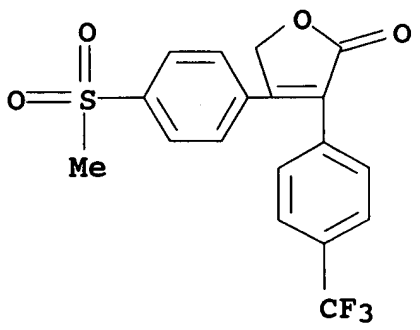
RN 162011-99-6 CAPLUS
CN 2(5H)-Furanone, 3-(2,6-dichlorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



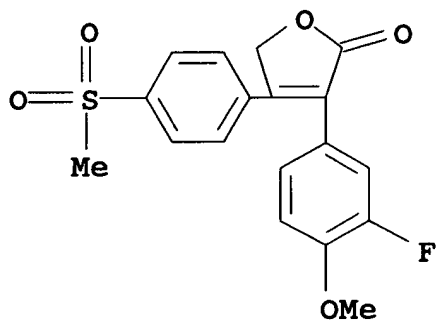
RN 162012-00-2 CAPLUS
CN 2(5H)-Furanone, 3-(3-chloro-4-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



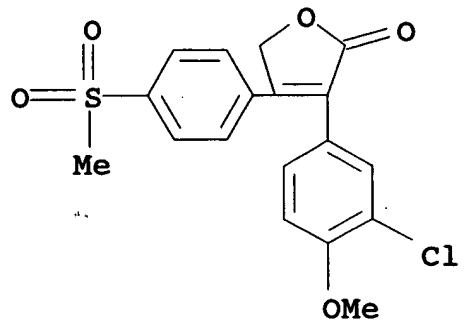
RN 162012-01-3 CAPLUS
CN 2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-[4-(trifluoromethyl)phenyl]- (9CI) (CA INDEX NAME)



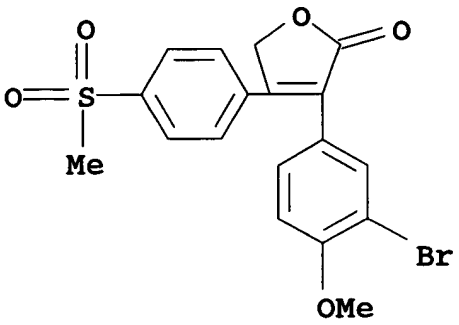
RN 162012-02-4 CAPLUS
CN 2(5H)-Furanone, 3-(3-fluoro-4-methoxyphenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



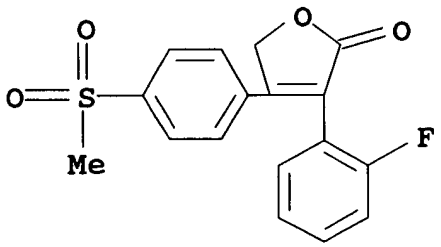
RN 162012-03-5 CAPLUS
CN 2(5H)-Furanone, 3-(3-chloro-4-methoxyphenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



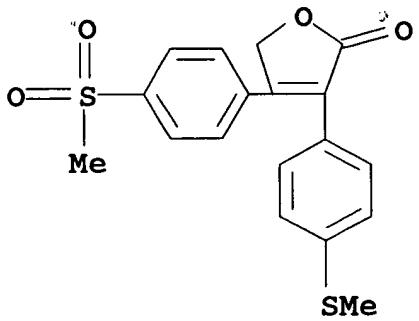
RN 162012-04-6 CAPLUS
CN 2(5H)-Furanone, 3-(3-bromo-4-methoxyphenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



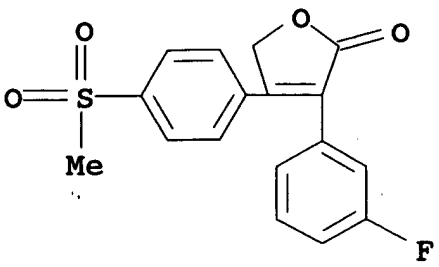
RN 162012-05-7 CAPLUS
CN 2(5H)-Furanone, 3-(2-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



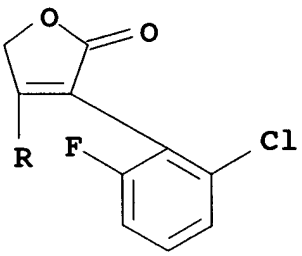
RN 162012-06-8 CAPLUS
CN 2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-[4-(methylthio)phenyl]- (9CI) (CA INDEX NAME)

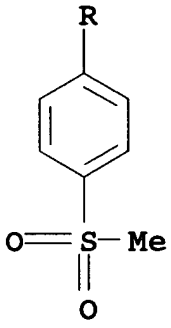


RN 162012-07-9 CAPLUS
CN 2(5H)-Furanone, 3-(3-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)

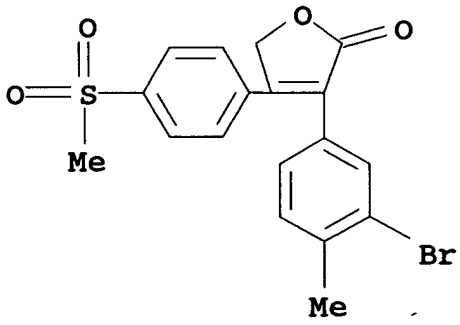


RN 162012-08-0 CAPLUS
CN 2(5H)-Furanone, 3-(2-chloro-6-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



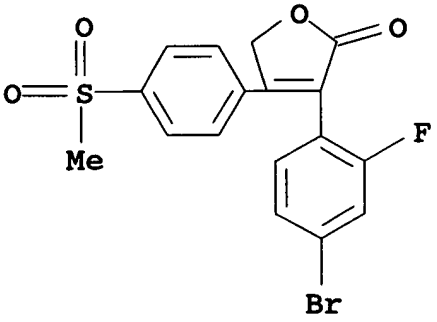


RN 162012-09-1 CAPLUS
CN 2(5H)-Furanone, 3-(3-bromo-4-methylphenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)

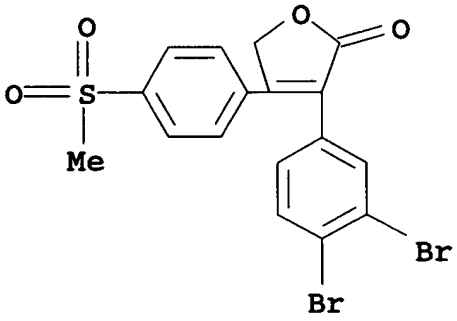


RN 162012-10-4 CAPLUS

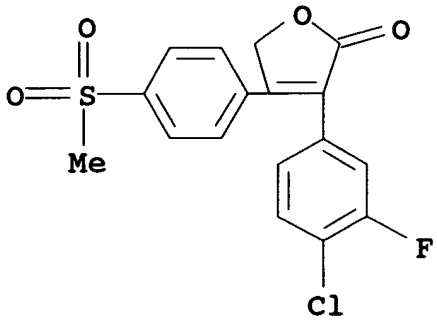
CN 2(5H)-Furanone, 3-(4-bromo-2-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



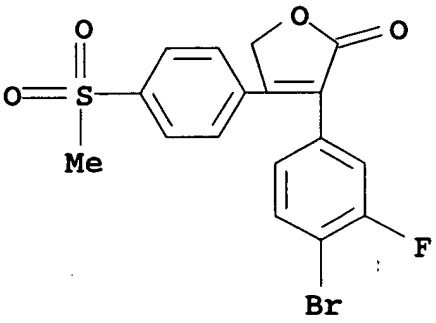
RN 162012-11-5 CAPLUS
CN 2(5H)-Furanone, 3-(3,4-dibromophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



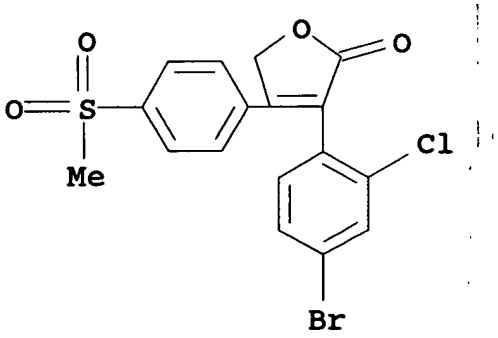
RN 162012-12-6 CAPLUS
CN 2(5H)-Furanone, 3-(4-chloro-3-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



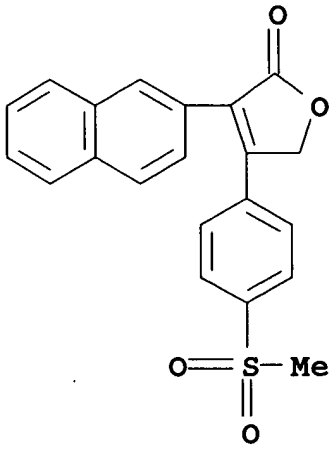
RN 162012-13-7 CAPLUS
CN 2(5H)-Furanone, 3-(4-bromo-3-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



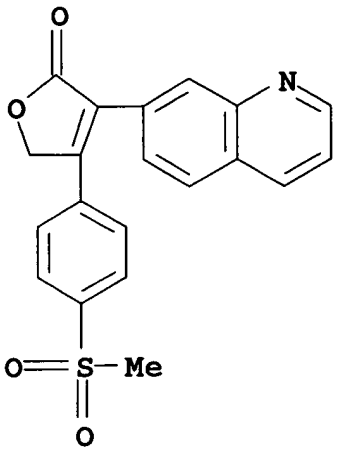
RN 162012-14-8 CAPLUS
CN 2(5H)-Furanone, 3-(4-bromo-2-chlorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



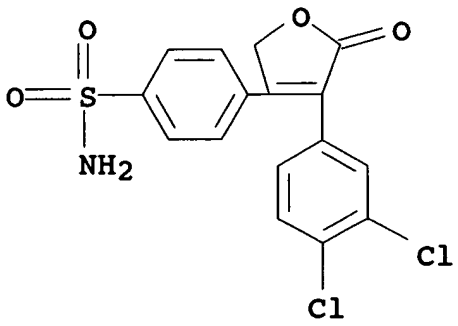
RN 162012-15-9 CAPLUS
CN 2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-(2-naphthalenyl)- (9CI) (CA INDEX NAME)



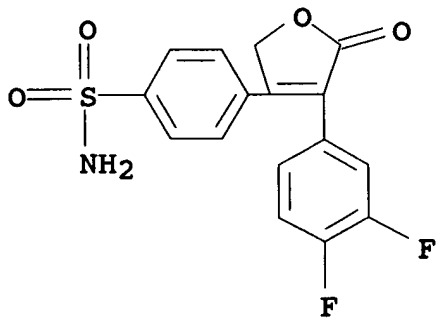
RN 162012-16-0 CAPLUS
CN 2(5H)-Furanone, 4-[4-(methylsulfonyl)phenyl]-3-(7-quinolinyl)- (9CI) (CA INDEX NAME)



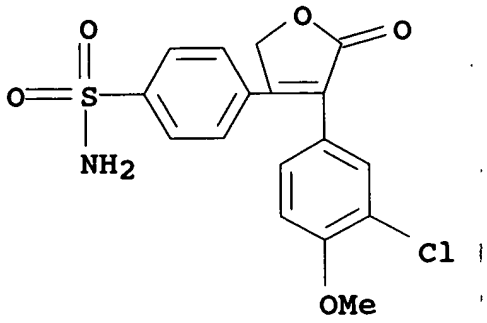
RN 162012-17-1 CAPLUS
CN Benzenesulfonamide, 4-[4-(3,4-dichlorophenyl)-2,5-dihydro-5-oxo-3-furanyl]- (9CI) (CA INDEX NAME)



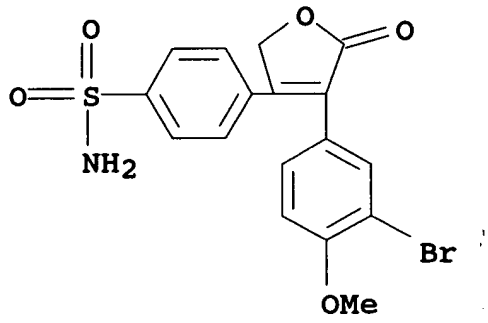
RN 162012-18-2 CAPLUS
CN Benzenesulfonamide, 4-[4-(3,4-difluorophenyl)-2,5-dihydro-5-oxo-3-furanyl]- (9CI) (CA INDEX NAME)



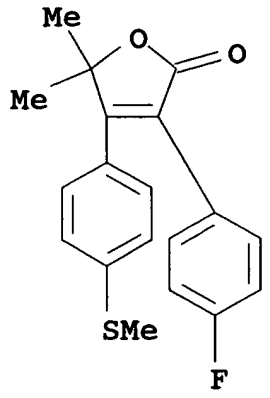
RN 162012-19-3 CAPLUS
CN Benzenesulfonamide, 4-[4-(3-chloro-4-methoxyphenyl)-2,5-dihydro-5-oxo-3-furanyl]- (9CI) (CA INDEX NAME)



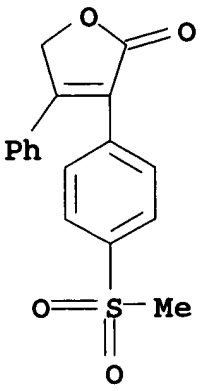
RN 162012-20-6 CAPLUS
CN Benzenesulfonamide, 4-[4-(3-bromo-4-methoxyphenyl)-2,5-dihydro-5-oxo-3-furanyl]- (9CI) (CA INDEX NAME)



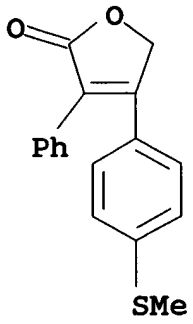
RN 162012-23-9 CAPLUS
CN 2(5H)-Furanone, 3-(4-fluorophenyl)-5,5-dimethyl-4-[4-(methylthio)phenyl]- (9CI) (CA INDEX NAME)



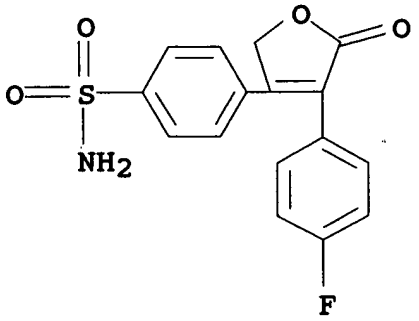
RN 162012-25-1 CAPLUS
CN 2(5H)-Furanone, 3-[4-(methylsulfonyl)phenyl]-4-phenyl- (9CI) (CA INDEX NAME)



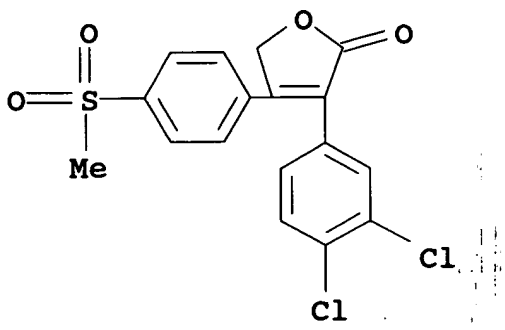
RN 162012-30-8 CAPLUS
CN 2(5H)-Furanone, 4-[4-(methylthio)phenyl]-3-phenyl- (9CI) (CA INDEX NAME)



RN 162012-33-1 CAPLUS
CN Benzenesulfonamide, 4-[4-(4-fluorophenyl)-2,5-dihydro-5-oxo-3-furanyl]- (9CI) (CA INDEX NAME)



RN 162012-35-3 CAPLUS
CN 2(5H)-Furanone, 3-(3,4-dichlorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)


O
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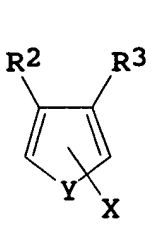
=> D BIB ABS HITSTR 3

L17 ANSWER 3 OF 5 CAPLUS COPYRIGHT 1995 ACS
AN 1994:579484 CAPLUS
DN 121:179484
TI Novel 3,4-diaryl thiophenes and analogs thereof having use as antiinflammatory agents
IN Bertenshaw, Stephen R.; Collins, Paul W.; Penning, Thomas D.; Reitz, David B.; Rogers, Roland S.
PA Searle, G. D., and Co., USA
SO PCT Int. Appl., 96 pp.
CODEN: PIXXD2
PI WO 9415932 A1 940721
DS W: AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB, HU, JP, KP, KR, KZ, LK, LU, LV, MG, MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD, SE, SK, UA, US, UZ, VN
RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, DE, DK, ES, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG
AI WO 94-US466 940114
PRAI US 93-4822 930115
DT Patent
LA English
OS MARPAT 121:179484
GI



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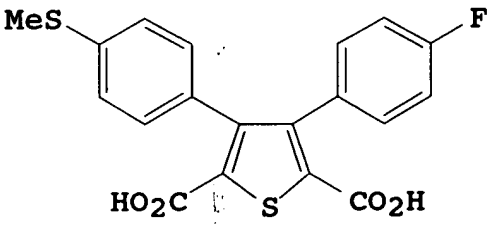
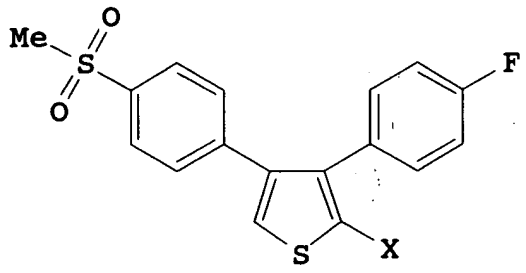


II



III

AB A class of 3,4-diaryl-substituted thiophene, furan, and pyrrole derivs. is disclosed, as well as pharmaceutical compns. contg. them, and methods of using them to treat inflammation and related disorders. Compds. of particular interest are I [Y = S, O, NR1; R1 = H, lower alkyl; X = 1 or 2 substituents selected from a large group, esp. H, halo, lower alkoxycarbonyl, CO2H; R2, R3 = (independently) aryl or heteroaryl, optionally substituted with 1 or more radicals such as sulfamyl, alkylsulfonyl, halo, lower alkoxy and lower alkyl] and pharmaceutically acceptable salts thereof. For example, S(CH2CO2Me)2 was cyclized with 4-FC6H4COCOC6H4(SMe)-4 in

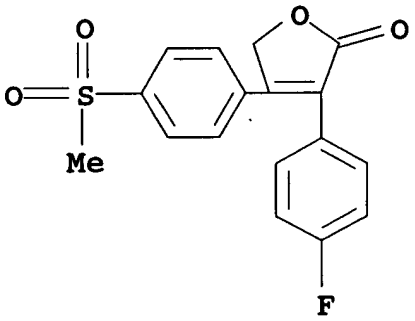
THF-MeOH contg. NaOMe at 65.degree. to give 82% of a mixt. of regioisomeric thiophenedicarboxylic acid monoesters, which were sapond. by NaOH in aq. THF-MeOH to give diacid II. Double decarboxylation of II with Cu in quinoline at 180-200.degree. (89%) and S-oxidn. with MCPBA gave title compd. III (X = H), which was brominated by Br2 in AcOH to give III (X = Br) plus the corresponding 2,5-dibromo compd. III (X = Br) at 10 mg/kg orally gave 30% inhibition in the carrageenan-induced rat-paw edema test. Data include 15 synthetic examples, rat-paw edema and analgesia tests, and in vitro tests for cyclooxygenase (I and II) and TXB2 activity.

IT **157672-00-9P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and reaction of, in prepn. of diarylthiophenes and analogs as antiinflammatory agents)

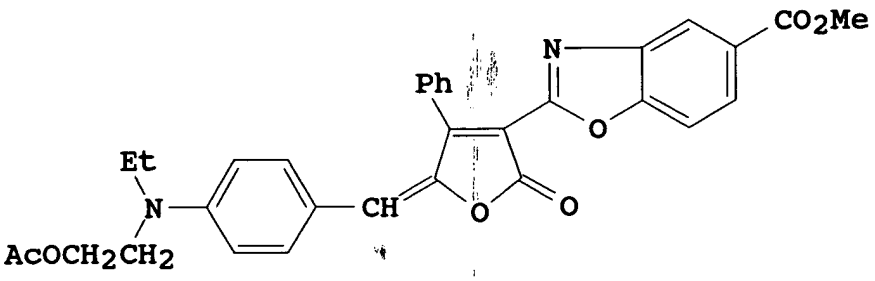
RN 157672-00-9 CAPLUS

CN 2(5H)-Furanone, 3-(4-fluorophenyl)-4-[4-(methylsulfonyl)phenyl]- (9CI) (CA INDEX NAME)



=> D BIB ABS HITSTR 4

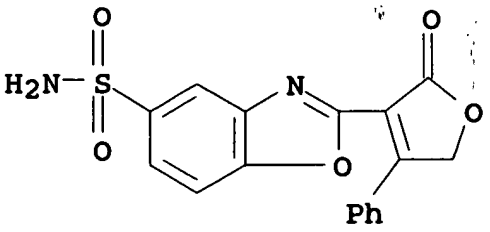
L17 ANSWER 4 OF 5 CAPLUS COPYRIGHT 1995 ACS
AN 1994:273062 CAPLUS
DN 120:273062
TI Polyesters having copolymerized therein a light-absorbing compound, and compositions thereof
IN Weaver, Max A.; Krutak, James J.; Coates, Clarence A., Jr.; Parham, William W.; Pruett, Wayne P.; Hilbert, Samuel D.
PA Eastman Kodak Co., USA
SO U.S., 45 pp.
CODEN: USXXAM
PI US 5274072 A 931228
AI US 92-878273 920504
DT Patent
LA English
GI



II

AB Substituted 2(5H)-furanones contg. a heteroaryl group in the 3-position and their derivs., useful as UV/visible light-absorbing compds., thermally stable with low volatility, were prepd. and copolymd. in high-temp. polyester prepns. or blended with polymers. Me 3-amino-4-hydroxybenzoate was cyclocondensed with EtO2CCH2C(OEt):NH.HCl to give Et (5-carbomethoxy-2-benzoxazolyl)acetate, which was cyclocondensed with PhCOCH2OAc to give 3-(5-carbomethoxy-2-benzoxazolyl)-4-phenyl-2(5H)-furanone (I). Condensation of I with 4-AcOCH2CH2NEtC6H4CHO gave II, .lambda.max 520 nm. Copolymn. of II 1.34 with 1,4-butanediol 81.0 and di-Me terephthalate 116.18 g gave a red polyester with wt.-av. mol. wt. 17,356 and polydispersity 1.40.

IT **154438-00-3P**
(prepn. of)
RN 154438-00-3 CAPLUS
CN 5-Benzoxazolesulfonamide, 2-(2,5-dihydro-2-oxo-4-phenyl-3-furanyl)- (9CI) (CA INDEX NAME)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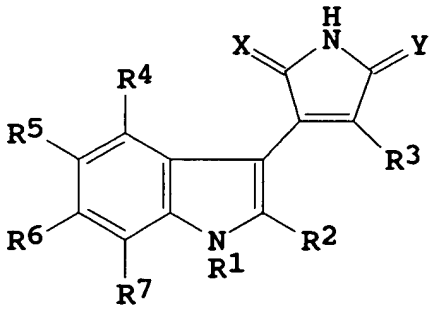
=> D BIB ABS HITSTR 5

L17 ANSWER 5 OF 5 CAPLUS COPYRIGHT 1995 ACS
AN 1990:98378 CAPLUS
DN 112:98378
TI Preparation of 3-(3-indolyl)pyrrole-2,5-diones and analogs as protein kinase inhibitors
IN Davis, Peter David; Hill, Christopher Huw; Lawton, Geoffrey
PA Hoffmann-La Roche, F., und Co. A.-G., Switz.
SO Eur. Pat. Appl., 38 pp.
CODEN: EPXXDW
PI EP 328026 A1 890816
DS R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
AI EP 89-102025 890206
PRAI GB 88-3048 880210
GB 88-27565 881125
DT Patent
LA German
GI



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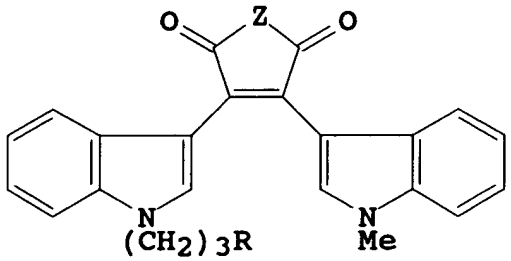
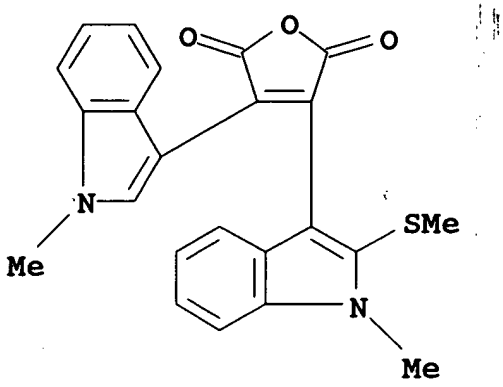


II

AB The title compds. (I; R1, R2 = H, alkyl, aryl, etc.; R3 = aryl, heteroaryl; R4-R7 = H, halo, alkyl, alkoxy, etc.; 1 of X, Y = O and the other = O, S, H and OH, H and H) were prepd. Thus, 1-(3-bromopropyl)indole (prepn. given) was stirred 2 h with (COCl)2 in CH2Cl2 and the product stirred 3 h with 1-methyl-3-indolylacetic acid in CH2Cl2 contg. (Me2CH)2NEt to give bis(indolyl)furandione II (R = Br, Z = O) which was converted in 3 steps to II (R = NH2, Z = NH). The latter was stirred 16 h with 1,1'-thiocarbonyldiimidazole in THF to give II (R = NCS, Z = NH) which had IC50 of 0.008 .mu.M for inhibition of protein kinase C in vitro.

IT **125314-93-4P**
(prepn. and reaction of, in prepn. of protein kinase inhibitors)
RN 125314-93-4 CAPLUS
CN 2,5-Furandione, 3-(1-methyl-1H-indol-3-yl)-4-[1-methyl-2-(methylthio)-1H-indol-3-yl]- (9CI) (CA INDEX NAME)


O
O
O
N
Me
SMe
N
Me

```
FILE 'CAOLD' ENTERED AT 11:09:35 ON 30 OCT 95
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 1995 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'MEDLINE' ENTERED AT 11:09:35 ON 30 OCT 95

FILE 'BIOSIS' ENTERED AT 11:09:35 ON 30 OCT 95
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=>
=>
=> D HIS

     (FILE 'CIN' ENTERED AT 10:27:46 ON 30 OCT 95)
                DEL HIS Y
                ACT JOHNMON/Q
               ---------
L1              STR
               ---------

     FILE 'REGISTRY' ENTERED AT 10:48:47 ON 30 OCT 95
L2            0 S L1
                E NC5/ESS(S)OC4/ESS(S)C6/ESS
L3            2 S L1 FUL

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     FILE 'CAPLUS' ENTERED AT 10:53:38 ON 30 OCT 95
L4            1 S L3

     FILE 'CAOLD, MEDLINE, BIOSIS' ENTERED AT 10:54:38 ON 30 OCT 95
L5            0 FILE CAOLD
L6            0 FILE MEDLINE
L7            0 FILE BIOSIS
     TOTAL FOR ALL FILES
L8            0 S L3

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     FILE 'HOME' ENTERED AT 10:55:35 ON 30 OCT 95

     FILE 'REGISTRY' ENTERED AT 11:00:13 ON 30 OCT 95
L9              STR
L10           0 S L9
L11             SCR 1840 AND 2005 AND 2021 AND 72
L12           7 S L9 AND L11
L13             STR L9
L14           5 S L9 AND L13 AND L11
L15          48 S L9 AND L13 AND L11 FUL

     FILE 'CAPLUS' ENTERED AT 11:08:09 ON 30 OCT 95
L16           5 S L15

     FILE 'REGISTRY' ENTERED AT 11:08:24 ON 30 OCT 95

     FILE 'CAPLUS' ENTERED AT 11:08:31 ON 30 OCT 95
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L17             5 S L15

     FILE 'CAOLD, MEDLINE, BIOSIS' ENTERED AT 11:09:35 ON 30 OCT 95
L18             0 FILE CAOLD
L19             0 FILE MEDLINE
L20             0 FILE BIOSIS
     TOTAL FOR ALL FILES
L21             0 S L15

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     FILE 'HCAPLUS' ENTERED AT 11:12:28 ON 30 OCT 95
L23             5 S L15
                  SELECT PRN L23 1-5
                  SELECT PN L23 1-5

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L24             0 S L22 NOT E1-10

=> FIL HOM
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